|                                                                                                              |                                                                                              |                                                                                                              |                                                       | A                                        |  |  |  |  |  |  |  |
|--------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------|-------------------------------------------------------|------------------------------------------|--|--|--|--|--|--|--|
| TO WELL DITTO OF E COMP SERVE                                                                                |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | TRANSMITTAL LETTER DESIGNATED/ELECTE                                                                         |                                                       | Attorney Docket No. 0512-1317            |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | CONCERNING A FILING                                                                                          |                                                       | U.S. Application No.                     |  |  |  |  |  |  |  |
| INT                                                                                                          | FRN                                                                                          | ATIONAL APPLN. NO.                                                                                           | INTERNATIONAL FILING DATE                             | 10/564,284 INTERNATIONAL APPLN. NO.      |  |  |  |  |  |  |  |
| PCT/IB2004/002265                                                                                            |                                                                                              |                                                                                                              | July 13, 2004                                         | PCT/IB2004/002265                        |  |  |  |  |  |  |  |
| TIT                                                                                                          | TITLE OF INVENTION: EXTRACELLULAR VESICLES FROM NON-PATHOGENIC AMOEBAE USEFUL AS VEHICLE FOR |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| TRASFERRING A MOLECULE OF INTEREST TO AN EUKARYOTIC CELL                                                     |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| APPLICANT(S) FOR DO/EO/US: Irène TATISCHEFF, Annette ALFSEN, Françoise LAVIALLE                              |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| Applicant herewith submits to the United States Designated Elected Office (DO/EO/US) the following items and |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| other information:                                                                                           |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| 1.                                                                                                           |                                                                                              |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| 2.                                                                                                           | $\boxtimes$                                                                                  | This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a filing under 35 U.S.C. 371.    |                                                       |                                          |  |  |  |  |  |  |  |
| 3.                                                                                                           |                                                                                              | This is an express request to begin national examination procedures (35 U.S.C. 371(f)).                      |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | The submission must include items (5), (6), (9) and (21) indicated below.                                    |                                                       |                                          |  |  |  |  |  |  |  |
| 4.                                                                                                           |                                                                                              | The US has been elected (Article 31).                                                                        |                                                       |                                          |  |  |  |  |  |  |  |
| 5.                                                                                                           |                                                                                              | A copy of the International Application as filed (35 U.S.C. 371 (c)(2))                                      |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | a.                                                                                           | is attached hereto (required only if not communicated by the International Bureau).                          |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | b.                                                                                           | has been communicated by the International Bureau. See attached PCT/IB/308.                                  |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | c.                                                                                           | is not required, as the application was filed in the United States Receiving Office (RO/US).                 |                                                       |                                          |  |  |  |  |  |  |  |
| 6.                                                                                                           |                                                                                              | An English language translation of the International Application as filed (35 U.S.C. 371 (c)(2))             |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | <u> —</u>                                                                                    | is attached hereto.                                                                                          | 11.                                                   | (=,(=,/,=,/,=,/,=,/,=,/,=,/,=,/,=,/,=,/, |  |  |  |  |  |  |  |
|                                                                                                              | b.                                                                                           | has been previously subm                                                                                     | nitted under 35 U.S.C. 154(d)(4).                     |                                          |  |  |  |  |  |  |  |
| 7.                                                                                                           |                                                                                              | Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371 (c)(3))        |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | a.                                                                                           | are attached hereto (required only if not communicated by the International Bureau).                         |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | b.                                                                                           | have been communicated by the International Bureau.                                                          |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | c.                                                                                           | have not been made, however, the time limit for making such amendments has NOT expired.                      |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | d.                                                                                           | have not been made, nowever, the time limit for making such amendments has NOT expired.                      |                                                       |                                          |  |  |  |  |  |  |  |
| 8.                                                                                                           |                                                                                              | An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371 (c)(3)). |                                                       |                                          |  |  |  |  |  |  |  |
| 9.                                                                                                           |                                                                                              | An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4))                                              |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | A Declaration of Inventorship for purposes of U.S.A. designation pursuant to rule 4.17(iv). ))               |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | An English language translation of the annexes of the International Preliminary Examination Report           |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | under PCT Article 36 (35 U.S.C. 371(c)(5)).                                                                  |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | Items 12 to 23 below concern document(s) or information included:                            |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| 12.                                                                                                          |                                                                                              |                                                                                                              | nent (IDS) w/PTO-1449 - 🔲 Copy of ID                  | S citations 🛛 PCT/ISA/210                |  |  |  |  |  |  |  |
| 13.                                                                                                          |                                                                                              | Assignment Papers (cover sh                                                                                  | eet & document(s)).                                   |                                          |  |  |  |  |  |  |  |
| 14.                                                                                                          |                                                                                              | A preliminary amendment.                                                                                     |                                                       |                                          |  |  |  |  |  |  |  |
| 15.                                                                                                          |                                                                                              | An Application Data Sheet under 37 C.F.R. 1.76.                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| 16.                                                                                                          |                                                                                              | Itemized Return Receipt Postcard                                                                             |                                                       |                                          |  |  |  |  |  |  |  |
| 17.                                                                                                          |                                                                                              | Abstract                                                                                                     |                                                       |                                          |  |  |  |  |  |  |  |
| 18.                                                                                                          |                                                                                              | A substitute specification.                                                                                  |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              |                                                                                              | •                                                                                                            | Power of Attorney and Statement under 37 CFR §3.73(b) |                                          |  |  |  |  |  |  |  |
|                                                                                                              | a. Newly executed Power of Attorney                                                          |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
|                                                                                                              | b. A change of Power of Attorney and/or change of address letter.                            |                                                                                                              |                                                       |                                          |  |  |  |  |  |  |  |
| 20.                                                                                                          |                                                                                              | _                                                                                                            | the sequence listing in accordance with               |                                          |  |  |  |  |  |  |  |

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22. A second copy of the English language translation of the international application (35 U.S.C. 154(d)(4)).

21. A second copy of the published international application under 35 U.S.C. 154(d)(4).

and 37 CFR 1.821 - 1.825.

23. Other items or information:

| U.S. APPLICATION NO.<br>10/564,284                                                                                                                                                                                     | INTERNATIONAL AF                                       |                                    |                   | EY DOCKET NO.<br>7 |                                 |  |  |  |  |  |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------|------------------------------------|-------------------|--------------------|---------------------------------|--|--|--|--|--|
| ☐ Applicant claims small er                                                                                                                                                                                            | Applicant claims small entity status. See 37 CFR 1.27. |                                    |                   |                    |                                 |  |  |  |  |  |
| ☐ The following fees are subr                                                                                                                                                                                          | The following fees are submitted:                      |                                    |                   |                    |                                 |  |  |  |  |  |
| PCT FEE                                                                                                                                                                                                                |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
|                                                                                                                                                                                                                        |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| <u>Fe</u>                                                                                                                                                                                                              |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
|                                                                                                                                                                                                                        |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| Non ISA/US Search Provided                                                                                                                                                                                             |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| US was the IPEA And all claims the provisions of PCT Article 33 (1)-(4                                                                                                                                                 |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| Surcharge of \$130.00 for furnishing the months from the earliest claimed priority                                                                                                                                     |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| National Stage Application                                                                                                                                                                                             | Additional Shee                                        |                                    | Fee From          |                    |                                 |  |  |  |  |  |
| size fee each additional 50<br>sheets in excess of 100                                                                                                                                                                 | - 100 = =                                              | X                                  | Below<br>\$125.00 | \$0.00             |                                 |  |  |  |  |  |
| Fee Code 1681/2681                                                                                                                                                                                                     | 50                                                     |                                    |                   |                    |                                 |  |  |  |  |  |
| CLAIMS                                                                                                                                                                                                                 | NUMBER FILED                                           | NUMBER EXTRA                       | RATE              |                    |                                 |  |  |  |  |  |
| Independent Claims Fee Codes 1614 / 2614 1 - 3 = 0 x \$100.00                                                                                                                                                          |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| Total Claims Fee Codes 1615                                                                                                                                                                                            | x \$25.00                                              | \$0.00                             |                   |                    |                                 |  |  |  |  |  |
| MULTIPLE DEPENDENT CLAIM(S) (if a                                                                                                                                                                                      | \$0.00                                                 |                                    |                   |                    |                                 |  |  |  |  |  |
| Processing fee of \$130.00 for furnishing months from the earliest claimed priority                                                                                                                                    | \$0.00                                                 |                                    |                   |                    |                                 |  |  |  |  |  |
| Fee for recording the enclosed assignm                                                                                                                                                                                 | \$0.00                                                 |                                    |                   |                    |                                 |  |  |  |  |  |
|                                                                                                                                                                                                                        |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
|                                                                                                                                                                                                                        | Amount to be refunded:                                 | \$                                 |                   |                    |                                 |  |  |  |  |  |
|                                                                                                                                                                                                                        |                                                        | ·                                  |                   | Charged:           | \$                              |  |  |  |  |  |
| A check in the amount of \$                                                                                                                                                                                            | to cover the above                                     | e fees is attached.                |                   |                    |                                 |  |  |  |  |  |
| The Director is hereby authorized to charge indicated fees and credit any overpayment to Deposit Account No. 25-0120 in the name of Young & Thompson. A duplicate copy of this sheet is enclosed.                      |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| The Director is hereby authorized in this, concurrent, and future replies, to charge payment or credit any overpayment to Deposit Account No. 25-0120 for any additional fee required under 37 C.F.R. §§ 1.16 or 1.17. |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| 1 26 6                                                                                                                                                                                                                 |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| SEND ALL CORRESPONDENCE TO YOUNG & THOMPSON                                                                                                                                                                            |                                                        | SIGNATURE                          |                   |                    |                                 |  |  |  |  |  |
| 745 South 23 <sup>rd</sup> Street<br>Arlington, VA 22202                                                                                                                                                               |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| Telephone: (703) 521-2297<br>Facsimile: (703) 685-0573                                                                                                                                                                 |                                                        | Benoit Castel, Re<br>NAME, REGISTF |                   |                    | eg. No. 35,041<br>RATION NUMBER |  |  |  |  |  |
| Y&T Customer No. <b>00466</b>                                                                                                                                                                                          |                                                        |                                    |                   |                    |                                 |  |  |  |  |  |
| BC/pmt                                                                                                                                                                                                                 |                                                        | March 14, 2006<br>DATE             |                   |                    |                                 |  |  |  |  |  |